

TABLE 2p

Representative compounds with activity toward kinase Yes with IC ₅₀ ≤10 μM under the test conditions employed.	
Yes:	P-0002

Example 13

Efficacy of Compounds in Combination with
Standard-of-Care Chemotherapeutic Agents in Four
Human Cancer Cell Lines

Compounds of the invention, such as compounds of Formula I, in combination with a standard chemotherapeutic agent, such as 5-fluorouracil, carboplatin, dacarbazine, gefitinib, oxaliplatin, paclitaxel, SN-38, temozolomide, or vinblastine, can be assessed for their effectiveness in killing human tumor cells. Such assays are known in the art, for example, as described in U.S. patent application Ser. No. 11/473,347.

Additional examples of certain methods contemplated by the present invention may be found in the following applications: U.S. Patent Publ. No. 2006/058339; U.S. Patent Publ. No. 2006/058340; U.S. Patent Publ. No. 2007/0032519; and U.S. patent application Ser. No. 11/473,347, filed Jun. 21, 2006 (Equivalent to PCT published as WO 2007/002433), each of which are hereby incorporated by reference herein in their entireties including all specifications, figures, and tables, and for all purposes.

All patents and other references cited in the specification are indicative of the level of skill of those skilled in the art to which the invention pertains, and are incorporated by reference in their entireties, including any tables and figures, to the same extent as if each reference had been incorporated by reference in its entirety individually.

One skilled in the art would readily appreciate that the present invention is well adapted to obtain the ends and advantages mentioned, as well as those inherent therein. The methods, variances, and compositions described herein as presently representative of preferred embodiments are exemplary and are not intended as limitations on the scope of the invention. Changes therein and other uses will occur to those skilled in the art, which are encompassed within the spirit of the invention, are defined by the scope of the claims.

The invention illustratively described herein suitably may be practiced in the absence of any element or elements, limitation or limitations which is not specifically disclosed herein. Thus, for example, in each instance herein any of the terms "comprising", "consisting essentially of" and "consisting of" may be replaced with either of the other two terms. Thus, for an embodiment of the invention using one of the terms, the invention also includes another embodiment wherein one of these terms is replaced with another of these terms. In each embodiment, the terms have their established meaning. Thus, for example, one embodiment may encompass a method "comprising" a series of steps, another embodiment would encompass a method "consisting essentially of" the same steps, and a third embodiment would encompass a method "consisting of" the same steps. The terms and expressions which have been employed are used as terms of description and not of limitation, and there is no intention that in the use of such terms and expressions of excluding any equivalents of the features shown and described or portions thereof, but it is recognized that

various modifications are possible within the scope of the invention claimed. Thus, it should be understood that although the present invention has been specifically disclosed by preferred embodiments and optional features, modification and variation of the concepts herein disclosed may be resorted to by those skilled in the art, and that such modifications and variations are considered to be within the scope of this invention as defined by the appended claims.

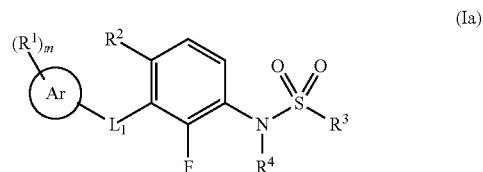
In addition, where features or aspects of the invention are described in terms of Markush groups or other grouping of alternatives, those skilled in the art will recognize that the invention is also thereby described in terms of any individual member or subgroup of members of the Markush group or other group.

Also, unless indicated to the contrary, where various numerical values are provided for embodiments, additional embodiments are described by taking any 2 different values as the endpoints of a range. Such ranges are also within the scope of the described invention.

Thus, additional embodiments are within the scope of the invention and within the following claims.

What is claimed is:

1. A compound of formula (Ia):



or a pharmaceutically acceptable salt thereof, wherein:

L₁ is a bond or —N(H)C(O)—;

each R¹ is optionally substituted lower alkyl or optionally substituted heteroaryl;

R² is hydrogen or halogen;

R⁴ is hydrogen;

R³ is optionally substituted lower alkyl or optionally substituted aryl;

m is 0, 1, 2, 3, 4, or 5; and

Ar is a monocyclic heteroaryl containing 5 to 6 atoms wherein at least one atom is nitrogen.

2. The compound of claim 1, wherein R² is hydrogen.

3. The compound of claim 1, wherein R² is halogen.

4. The compound of claim 1, wherein R³ is optionally substituted phenyl.

5. The compound of claim 1, wherein R³ is phenyl substituted with one or more substituents selected from the group consisting of fluoro, lower alkyl, fluoro substituted lower alkyl, lower alkoxy, fluoro substituted lower alkoxy, lower alkylthio, and fluoro substituted lower alkylthio.

6. The compound of claim 1, wherein R³ is phenyl substituted with one or more fluoro.

7. The compound of claim 1, wherein each R¹ is optionally substituted lower alkyl.

8. The compound of claim 1, wherein each R¹ is optionally substituted heteroaryl.

9. A pharmaceutical composition comprising a compound of claim 1 and a pharmaceutically acceptable carrier or excipient.

10. A pharmaceutical composition comprising a compound of claim 1 and another therapeutic agent.